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or proteinase or enzyme (2A) activity)
L1         191 ((KLK 8) OR (KALLIKREIN 8) OR NEUROPSIN OR OVASIN) (6A) (PROTEAS
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OR EVALUATED OR EVALUATION)
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L3       38846 (PROTEASE OR PEPTIDASE OR PROTEINASE OR ENZYME (2A) ACTIVITY)
               (6A) (METHOD OR ANALYSIS OR ANALYZE OR ANALYZED OR ANALYZING OR
               TEST OR TESTING OR TESTED OR DETERMINING OR DETERMINATION OR
               DETERMINE OR DETERMINED OR EVALUATE OR EVALUATION OR EVALUATED
               OR EVALUATION)

=> s l2 and l3
L4           6 L2 AND L3

=> d l5 1-6 bib ab
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=> d l4 1-6 bib ab

L4   ANSWER 1 OF 6     MEDLINE on STN
AN   97312794     MEDLINE
DN   PubMed ID: 9169235
TI   Crystallization and preliminary X-ray analysis of
     neuropsin, a serine protease expressed in the limbic
     system of mouse brain.
AU   Kishi T; Kato M; Shimizu T; Kato K; Matsumoto K; Yoshida S; Shiosaka S;
     Hakoshima T
CS   Department of Molecular Biology, Nara Institute of Science and Technology
     (NAIST), Japan.
SO   Journal of structural biology, (1997 Apr) Vol. 118, No. 3, pp. 248-51.
     Journal code: 9011206. ISSN: 1047-8477.
CY   United States
DT   Journal; Article; (JOURNAL ARTICLE)
     (RESEARCH SUPPORT, NON-U.S. GOV'T)
LA   English
FS   Priority Journals
EM   199706
ED   Entered STN: 9 Jul 1997
     Last Updated on STN: 3 Mar 2000
     Entered Medline: 20 Jun 1997
AB   Neuropsin (M(r) 25032) is a serine protease expressed
     in the limbic system of mouse brain.  It has been implicated in various
     neurological processes including formation of memory and may be important
     as a drug target in the treatment of epilepsy.  The recombinant protein
     was produced using a baculovirus expression system and was purified.  Two
     crystal forms were obtained by a hanging-drop vapor-diffusion method with
     polyethylene glycol.  Preliminary X-ray crystallographic analysis revealed
     that crystal form I belongs to triclinic space group P1 with unit cell
     dimensions a = 97.16 A, b = 97.12 A, c = 46.75 A and alpha = 99.17
     degrees, beta = 99.77 degrees, gamma = 117.35 degrees.  Self-rotation
     function analysis of these data of form I indicates the position of a
     noncrystallographic threefold axis.  There are six molecules in the
     crystallographic asymmetric unit.  Crystal form II also belongs to
     triclinic space group P1 but has unit cell dimensions of a = 38.40 A, b =
     55.16 A, c = 65.37 A and alpha = 95.38 degrees, beta = 89.98 degrees,
     gamma = 110.46 degrees with two molecules in the crystallographic
     asymmetric unit.  Form II has a noncrystallographic twofold axis.
     Intensity data to 3.1 A resolution for form I and to 2.2 A resolution for
     form II have been collected.

L4   ANSWER 2 OF 6  BIOSIS  COPYRIGHT (c) 2008 The Thomson Corporation  on STN
AN   2006:398347  BIOSIS
DN   PREV200600398661
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TI Methods and reagents for protease inhibition.
AU Albrecht, Hugo [Inventor]; Hengst, Ulrich [Inventor]; Monard, Denis [Inventor]
CS Riehen, Switzerland
ASSIGNEE: Novartis Forschungsstiftung Zweigniederlassung Friedrich Miescher Instittue for Biomedical Research
PI US 07029877 20060418
SO Official Gazette of the United States Patent and Trademark Office Patents, (APR 18 2006)
CODEN: OGUPE7. ISSN: 0098-1133.
DT Patent
LA English
ED Entered STN: 9 Aug 2006
Last Updated on STN: 9 Aug 2006
AB There is provided a protease inhibitor and a method of inhibiting a protease selected from the group consisting of thrombin, chymotrypsin and neuropsin, by contacting the protease with an effective amount of a member of the phosphoethanolamine binding protein (PEBP) family.

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
AN 2006:1248277 CAPLUS
DN 146:22551
TI Random mutagenesis, screening and selection of protease variants with altered sensitivity to activity modulators
IN Koltermann, Andre; Kettling, Ulrich; Haupts, Ulrich; Coco, Wayne; Tebbe, Jan; Votsmeier, Christian; Scheidig, Andreas
PA Direvo Biotech AG, Germany
SO Eur. Pat. Appl., 93pp.
CODEN: EPXXDW
DT Patent
LA English
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 1726643	A1	20061129	EP 2005-104543	20050527
	R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, LV, MK, YU				
	US 20060269538	A1	20061130	US 2006-441635	20060526
	WO 2006125827	A1	20061130	WO 2006-EP62644	20060526
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	EP 1883696	A1	20080206	EP 2006-763303	20060526
	R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR				
PRAI	EP 2005-104543	A	20050527		
	US 2005-685566P	P	20050527		
	US 2005-686021P	P	20050531		
	WO 2006-EP62644	W	20060526		

AB The present invention provides a method for the selection of proteases

with altered sensitivity to one or more activity-modulating substances. The method combines the provision of a protease library (i.e., phage display library) encoding polynucleotide sequences generated by using PCR mutagenesis, expression of the enzymes, screening of the library in the presence of one or several activity-modulating substances, selection of variants with altered sensitivity to one or several activity-modulating substances and isolation of those polynucleotide sequences that encode for the selected variants. In particular, mutant variants of human trypsin are disclosed.

RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
AN 2006:684183 CAPLUS
DN 146:2865
TI Activation and enzymatic characterization of recombinant human kallikrein 8
AU Kishi, Tadaaki; Cloutier, Sylvain M.; Kundig, Christoph; Deperthes, David; Diamandis, Eleftherios P.
CS Department of Pathology and Laboratory Medicine, Mount Sinai Hospital, Toronto, ON, M5G 1X5, Can.
SO Biological Chemistry (2006), 387(6), 723-731
CODEN: BICHF3; ISSN: 1431-6730
PB Walter de Gruyter GmbH & Co. KG
DT Journal
LA English
AB Human kallikrein 8 (hK8), whose gene was originally cloned as the human ortholog of a mouse brain protease, is known to be associated with diseases such as ovarian cancer and Alzheimer's disease. Recombinant human pro-kallikrein 8 was activated with lysyl endopeptidase-conjugated beads. Amino-terminal sequencing of the activated enzyme demonstrated the cleavage of a 9-aa propeptide from the pro-enzyme. The substrate specificity of activated hK8 was characterized using synthetic fluorescent substrates. HK8 showed trypsin-like specificity, as predicted from sequence anal. and enzymic characterization of the mouse ortholog. All synthetic substrates tested containing either arginine or lysine at P1 position were cleaved by hK8. The highest kcat/Km value of 20+103 M-1 s-1 was observed with Boc-Val-Pro-Arg-7-amido-4-methylcoumarin. The activity of hK8 was inhibited by antipain, chymostatin, and leupeptin. The concentration for 50% inhibition by the best inhibitor, antipain, was 0.46 μM. The effect of different metal ions on the enzyme activity was analyzed. Whereas Na+ had no effect on hK8 activity, Ni2+ and Zn2+ decreased the activity and Ca2+, Mg2+, and K+ had a stimulatory effect. Ca2+ was the best activator, with an optimal concentration of approx. 10 μM.

RE.CNT 61 THERE ARE 61 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
AN 2005:1020554 CAPLUS
DN 143:282218
TI Protease activity assay method by using polymeric membrane
IN Shiosaka, Sadao; Tamura, Hidenori
PA Nara Institute of Science and Technology, Japan
SO Jpn. Kokai Tokkyo Koho, 17 pp.
CODEN: JKXXAF
DT Patent
LA Japanese
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 2005253436	A	20050922	JP 2004-73625	20040315
PRAI	JP 2004-73625		20040315		

AB A assay method for measuring protease (especially, neuropsin) activity with higher sensitivity and fewer sample amount than the traditional solution method. The method includes processes of (1) the sample containing protease is sticked to a polymeric membrane, (2) the protease is reacted with the substrate specific to the protease and (3) the signal resulted from the reaction is measured.

L4 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
AN 2002:172125 CAPLUS
DN 136:212778
TI Identification of a novel brain serine protease inhibitory protein-phosphoethanolamine binding protein and methods and reagents for protease inhibition for the treatment of neurological disorders
IN Albrecht, Hugo; Hengst, Ulrich; Monard, Denis
PA Novartis Forschungsstiftung Zweigniederlassung Friedrich Miescher Institute for Biomedical Research, Switz.
SO PCT Int. Appl., 39 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002018623	A2	20020307	WO 2001-EP10043	20010830
	WO 2002018623	A3	20021114		
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW				
	RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	CA 2420832	A1	20020307	CA 2001-2420832	20010830
	AU 2002012184	A	20020313	AU 2002-12184	20010830
	EP 1315758	A2	20030604	EP 2001-980309	20010830
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
	JP 2004507471	T	20040311	JP 2002-522529	20010830
	US 20050037009	A1	20050217	US 2003-362642	20030224
	US 7029877	B2	20060418		
	US 20060177432	A1	20060810	US 2005-311974	20051219
PRAI	GB 2000-21497	A	20000901		
	WO 2001-EP10043	W	20010830		
	US 2003-362642	A1	20030224		

AB The present invention is based on the discovery of a novel serine protease inhibitory protein-phosphoethanolamine binding protein (PEBP). PEPB is identified by the detection of a novel thrombin inhibitory activity in the brain of protease nexin-1(-/-) mice, a gene knockout for the only known endogenous protease inhibitor protease nexin-1 that specifically interferes with thrombotic activity and is expressed in the brain. PEBP exerts inhibitory activity against several serine proteases including thrombin, neuropsin, and chymotrypsin, whereas trypsin, tissue type plasminogen activator, and elastase are not affected. PEBP immunoreactivity is found on the surface of Rat-1 fibroblast cells and although its sequence contains no secretion signal, PEBP-H6 can be

purified from the conditioned medium upon recombinant expression. The method of inhibiting a protease selected from the group consisting of thrombin, chymotrypsin and neuropsin, by contacting the protease with an effective amount of PEBP.